0			2002/07/0 2 17:14	USPAT; US-PGPUB; EPO; JPO; DERWENT	1 same 8	0	Г9	BRS	9
0			2002/07/0 2 17:14	USPAT; US-PGPUB; EPO; JPO; DERWENT	((human adj growth adj hormone) or hgh ) same microsphere	5 1	L8	BRS	ω
0			2002/07/0 2 17:14	USPAT; US-PGPUB; EPO; JPO; DERWENT	19300 microsphere	19300	L7	BRS	7
0			2002/07/0 2 17:12	USPAT; US-PGPUB; EPO; JPO; DERWENT	((((human adj growth adj hormone) or hgh) same replacement) same recombinant) same dose	Н	Г6	BRS	δ
0			2002/07/0 2 17:11	USPAT; US-PGPUB; EPO; JPO; DERWENT	((human adj growth adj hormone) or hgh) same replacement	132	Ľ5	BRS	σ
0			2002/07/0 2 17:11	USPAT; US-PGPUB; EPO; JPO; DERWENT	((human adj growth adj hormone) or hgh) same (replensh or replenshing)	0	<b>Г</b> 4	BRS	4
0			2002/07/0 2 16:51	USPAT; US-PGPUB; EPO; JPO; DERWENT	1 same 2	16	ĽЗ	BRS	ω
0			2002/07/0 2 16:51	USPAT; US-PGPUB; EPO; JPO; DERWENT	(human adj growth adj hormone) or hgh	5314	L2	BRS	Ν
0			2002/07/0 2 16:50	USPAT; US-PGPUB; EPO; JPO; DERWENT	replacement adj therapy	3120	Ľ	BRS	Ľ
r a H	Erro r Defi niti on	Com men	Time Stamp	DBs	Search Text	Hits	۲ #	Туре	

=> d his

## (FILE 'HOME' ENTERED AT 17:17:25 ON 02 JUL 2002)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT

17:18:01 ON 02 JUL 2002

- L1 36627 S (HUMAN GROWTH HORMONE) OR HGH
- L2 92997 S (REPLACEMENT THERAPY) OR REPLENISH?
- L3 456 S L1 (P) L2
- L4 122 S L3 (P) DOSE
- L5 37 S L4 (P) DAILY
- L6 12 DUPLICATE REMOVE L5 (25 DUPLICATES REMOVED)
- L7 75372 S MICROSPHERE
- L8 144 S L1 (P) L7
- L9 0 S L8 (P) L2

 $\Rightarrow$  log y

FILE 'HOME' ENTERED AT 17:17:25 C JUL 2002 => file medline caplus biosis embase scisearch agricola TOTAL. COST IN U.S. DOLLARS SINCE FILE ENTRY SESSION 0.21 0.21 FULL ESTIMATED COST FILE 'MEDLINE' ENTERED AT 17:18:01 ON 02 JUL 2002 FILE 'CAPLUS' ENTERED AT 17:18:01 ON 02 JUL 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'BIOSIS' ENTERED AT 17:18:01 ON 02 JUL 2002 COPYRIGHT (C) 2002 BIOLOGICAL ABSTRACTS INC. (R) FILE 'EMBASE' ENTERED AT 17:18:01 ON 02 JUL 2002 COPYRIGHT (C) 2002 Elsevier Science B.V. All rights reserved. FILE 'SCISEARCH' ENTERED AT 17:18:01 ON 02 JUL 2002 COPYRIGHT (C) 2002 Institute for Scientific Information (ISI) (R) FILE 'AGRICOLA' ENTERED AT 17:18:01 ON 02 JUL 2002 => s (human growth hormone) or hgh 4 FILES SEARCHED... 36627 (HUMAN GROWTH HORMONE) OR HGH => s (replacement therapy) or replenish? 92997 (REPLACEMENT THERAPY) OR REPLENISH? => s 11 (p) 12456 L1 (P) L2 => s 13 (p) dose 122 L3 (P) DOSE => s 14 (p) daily 37 L4 (P) DAILY => duplicate remove 15 DUPLICATE PREFERENCE IS 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH' KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n PROCESSING COMPLETED FOR L5 12 DUPLICATE REMOVE L5 (25 DUPLICATES REMOVED) => d 16 1-12 ibib abs ANSWER 1 OF 12 MEDLINE DUPLICATE 1 ACCESSION NUMBER: 1998412518 MEDLINE 98412518 PubMed ID: 9741487 DOCUMENT NUMBER: TITLE: Periodic growth in rats. AUTHOR: Rol de Lama M A; Perez-Romero A; Ariznavarreta M C; Hermanussen M; Tresquerres J A Medical School, University Complutense, Madrid, Spain. CORPORATE SOURCE: ANNALS OF HUMAN BIOLOGY, (1998 Sep-Oct) 25 (5) 441-51. SOURCE: Journal code: 0404024. ISSN: 0301-4460. PUB. COUNTRY: ENGLAND: United Kingdom Journal; Article; (JOURNAL ARTICLE) LANGUAGE: English FILE SEGMENT: Priority Journals ENTRY MONTH: 199901 ENTRY DATE: Entered STN: 19990202 Last Updated on STN: 19990202 Entered Medline: 19990115 AB Microknemometry, a novel non-invasive technique, allows the accurate measurements of the lower leg length in the conscious rat, not only but even in periods smaller than 24 hours. Its use revealed the presence of nonlinear growth increments (mini-growth spurts) with

periods between 4 and 5 days, that presented a gradual decline in

```
***Hormone***
                                                                    (rhGH)
                  ***human***
                                  ***Growth***
     stimulated growth velocity in female rates, but did not show any effect on
     males. Neonatal Monosodium Glutamate (MSG) treatment reduced growth both
     in males and females. Growth hormone (GH) ***replacement***
                     in MSG treated animals was capable of increasing growth
     velocity, from day 30 onwards. The recovery was partial in males and
     complete in females. In intact male rats growth blockade induced by
     fasting was not followed by a catch up effect after refeeding, although
     growth velocity tended to increase and a clear catch up effect on weight
     was detected. Male rats seemed to grow at a maximal speed over at least
     the first 60 days of life, that cannot be accelerated with GH treatment,
     whereas female rats did respond to exogenous GH.
     ANSWER 2 OF 12 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
ACCESSION NUMBER:
                   1997:356620 BIOSIS
DOCUMENT NUMBER:
                    PREV199799663023
                    Growth hormone treatment for growth hormone deficient
TITLE:
                    adults.
AUTHOR (S):
                    Clark, W. (1); Kendall, M. J.
                    (1) Dep. Med. Management, Keele Univ., Keele, Staffordshire
CORPORATE SOURCE:
                    ST5 5BG UK
                    Journal of Clinical Pharmacy and Therapeutics, (1997) Vol.
SOURCE:
                    21, No. 6, pp. 367-372.
                    ISSN: 0269-4727.
                    Journal; Article
DOCUMENT TYPE:
                    English
LANGUAGE:
     Growth hormone (GH) deficiency in adults is now recognized as a clinical
     syndrome with characteristic signs and symptoms. Numerous trials with
                                               ***human***
       ***daily***
                     subcutaneous biosynthetic
       ***hormone*** ( ***hGH*** ) have been conducted in this patient group.
     Generally, improvements in insulin-like growth factor levels, decreases in
     total fat mass and increases in lean body mass are recorded with no
     overall effect on total body weight. Variable effects on serum
     cholesterol, bone mineral density and quality of life have also been
                                     ***replacement***
     reported. The true place of GH
                                                            ***therapy***
     adults has yet to be defined. Several questions relating to the
       ***dose*** , duration of treatment, long-term side-effects, quality of
     life changes and health economic implications of treatment still need to
     be assessed.
     ANSWER 3 OF 12 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE
     2
ACCESSION NUMBER:
                    1997:356635 BIOSIS
                    PREV199799663038
DOCUMENT NUMBER:
                    Effects of growth hormone replacement therapy on glucose
TITLE:
                    metabolism are due to changes of body composition.
                    Feldmeier, Horst O. (1); Nass, Ralf M.; Landgraf, Ruediger;
AUTHOR(S):
                    Strasburger, Christian J.
                    (1) Endokrinologische Abteilung, Med. Klinik, Klinikum
CORPORATE SOURCE:
                    Innenstadt, Ziemssenstrasse 1, 80366 Muenchen Germany
                    Journal of Pediatric Endocrinology & Metabolism, (1997)
SOURCE:
                    Vol. 10, No. SUPPL. 1, pp. 151-159.
DOCUMENT TYPE:
                    Article
LANGUAGE:
                    English
     The effects of 32 months
                                                      ***therapy***
                                ***replacement***
                                                                      with
                                  ***growth***
                                                    ***hormone***
                  ***human***
     recombinant
       ***hGH*** ) on glucose metabolism and body composition were investigated
     in 14 non diabetic, obese adult patients out of 21 patients with adult
     onset GH deficiency who were treated in this clinical trial. The
                                   ***dose***
                     ***hGH***
                                                  (12.5 pig/kg body weight) was
       ***daily***
     self administered subcutaneously. Oral glucose tolerance tests (OGTT) and
     measurement of IGF-I and HbA-1c were performed at the start and after 6,
                                        ***replacement***
                          ***hGH***
                                                               ***therapy***
     18 and 32 months of
     Body composition was evaluated by potassium-40 measurement at the start of
     the study, and after 6 and 18 months. For statistical analysis the
     Wilcoxon signed rank test and a linear, simple regression analysis were
                                                      ***hGH*** replacement
     performed; the results are given as mean +- SE.
```

significantly increased IGF-I levels (70.5+-9.5 ng/ml vs 264.7 +- 27.6

L6

amplitude when the animals were getting older, and a maximal growth rate between 0600h and 0900h. A small dimorphic growth pattern complete established with females growing less and presenting spurts of lower amplitude and smaller duration than males. High \*\*\*doses\*\*\* of

ng/ml; p lt 0.002). There was a positive correlation between the

\*\*\*hGH\*\*\*

\*\*\*dose\*\*\*

In the IGF-I levels (R =0.62, p 0.0001).

The HbA-1c levels significantly and constantly decreased during the 18 and
32 months of growth hormone replacement (5.4 +- 0.1 vs 4.7 +- 0.1%; p lt
0.002). The area under the curve (AUC) of the insulin values during the
OGTT decreased significantly after 18 and 32 months of \*\*\*hGH\*\*\*

replacement (16.0 +- 3.6 vs 10.6 +- 1.9 U/ml times 120 min and 12.2+-2
U/ml times 120 min; p lt 0.05). The lean body mass increased (49.7 +- 7.1
vs 53.7 +- 7.8 kg; p lt 0.002) and the fat mass significantly decreased
(39.0 +- 11.2 vs 35.4 +- 9.2 kg; p lt 0.002) during 18 months of

\*\*\*hGH\*\*\*

replacement. We observed a positive correlation between the
AUC of the insulin values and the fat mass (R=0.5; p lt 0.001).

\*\*\*hGH\*\*\*

replacement induces an initial insulin antagonistic effect,
followed by an apparent improvement in glucose utilization resulting from
a decrease of fat mass and increase of lean body mass.

L6 ANSWER 4 OF 12 MEDLINE DUPLICATE 3

ACCESSION NUMBER: 97345195 MEDLINE

DOCUMENT NUMBER: 97345195 PubMed ID: 9201562

TITLE: Growth hormone treatment for growth hormone deficient

adults.

AUTHOR: Clark W; Kendall M J

CORPORATE SOURCE: Department of Medicines Management, Keele University, U.K.

SOURCE: JOURNAL OF CLINICAL PHARMACY AND THERAPEUTICS, (1996 Dec)

21 (6) 367-72. Ref: 24

Journal code: 8704308. ISSN: 0269-4727.

PUB. COUNTRY: ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199708

ENTRY DATE: Entered STN: 19970813

Last Updated on STN: 19970813 Entered Medline: 19970801

AB Growth hormone (GH) deficiency in adults is now recognized as a clinical syndrome with characteristic signs and symptoms. Numerous trials with \*\*\*daily\*\*\* subcutaneous biosynthetic \*\*\*human\*\*\* \*\*\*growth\*\*\* \*\*\*hormone\*\*\* ( \*\*\*hGH\*\*\* ) have been conducted in this patient group. Generally, improvements in insulin-like growth factor levels, decreases in total fat mass and increases in lean body mass are recorded with no overall effect on total body weight. Variable effects on serum cholesterol, bone mineral density and quality of life have also been reported. The true place of GH \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* in adults has yet to be defined. Several questions relating to the

\*\*\*dose\*\*\* , duration of treatment, long-term side-effects, quality of life changes and health economic implications of treatment still need to be assessed.

L6 ANSWER 5 OF 12 MEDLINE DUPLICATE 4

ACCESSION NUMBER: 95155540 MEDLINE

DOCUMENT NUMBER: 95155540 PubMed ID: 7852519

TITLE: Effect of growth hormone (hGH) replacement therapy on

physical work capacity and cardiac and pulmonary function in patients with hGH deficiency acquired in adulthood.

AUTHOR: Nass R; Huber R M; Klauss V; Muller O A; Schopohl J;

Strasburger C J

CORPORATE SOURCE: Medical Clinic, Innenstadt University Hospital,

Ludwig-Maximilians-Universitat, Munich, Germany.

SOURCE: JOURNAL OF CLINICAL ENDOCRINOLOGY AND METABOLISM, (1995

Feb) 80 (2) 552-7.

Journal code: 0375362. ISSN: 0021-972X.

PUB. COUNTRY: United States

(CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199503

ENTRY DATE: Entered STN: 19950322

Last Updated on STN: 19950322

Entered Medline: 19950310

The effects of 6 months of \*replacement\*\*\* \*\*\*therapy with recombinant human GH ( \*\*\*hGH\*\*\* ) on physical work capacity and cardiac structure and function were investigated in 20 patients with \*\*\*hGH\*\*\* deficiency of adult onset in a double blind, placebo-controlled trial. The GH \*\*\*dose\*\*\* of 12.5 micrograms/kg BW was self-administered \*\*\*daily\*\*\* sc. Oxygen consumption (VO2), CO2 production, and ventilatory volumes were measured during exercise on a bicycle spiroergometer. M-Mode echocardiography was performed using standard

techniques. The VO2 max data, expressed per kg BW (mL/min.kg BW) showed a significant increase from 23.2  $\pm$ /- 2.4 to 30.0  $\pm$ /- 2.3 (P < 0.01) in the -treated group, whereas the VO2 max data, expressed per lean \*\*\*hGH\*\*\* body mass (milliliters per min/kg lean body mass) did not change significantly in either group. Maximal O2 pulse (milliliters per beat) increased significantly from 15.2 +/- 5.6 to 19.6 +/- 3.3 mL/beat (P < 0.01), but remained constant in the placebo group. The maximal power output (watts +/- SE) increased significantly (P < 0.01) from 192.5 +/-13.5 to 227.5 +/- 11.5 in the \*\*\*hGH\*\*\* -treated group, but remained constant in the placebo group. Cardiac structure (left ventricular posterior wall, interventricular septum thickness, left ventricular mass, left ventricular end-systolic dimension, and left ventricular end-diastolic dimension) as well as echocardiographically assessed cardiac function did not change significantly after 6 months of treatment in either group. We conclude that \*\*\*hGH\*\*\* replacement in -deficient adults improves oxygen uptake and exercise capacity. These improvements in pulmonary parameters might be due to an increase in respiratory muscle strength and partly to the changes in muscle volume per \*\*\*hGH\*\*\* \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* se observed during Furthermore, an increased cardiac output might contribute to the improvement in exercise performance during \*\*\*hGH\*\*\* treatment. \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* \*\*\*hGH\*\*\* According to our data, leads to an improvement of exercise capacity and maximal oxygen uptake, but has no significant effect on cardiac structure.

L6 ANSWER 6 OF 12 MEDLINE DUPLICATE 5

ACCESSION NUMBER: 95131154 MEDLINE

DOCUMENT NUMBER: 95131154 PubMed ID: 7830028

TITLE: Cardiovascular effects of prolonged growth hormone

replacement in adults.

AUTHOR: Beshyah S A; Shahi M; Foale R; Johnston D G

CORPORATE SOURCE: Unit of Metabolic Medicine, St Mary's Hospital and Medical

School, London, UK.

SOURCE: JOURNAL OF INTERNAL MEDICINE, (1995 Jan) 237 (1) 35-42.

Journal code: 8904841. ISSN: 0954-6820.

PUB. COUNTRY: ENGLAND: United Kingdom

(CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199502

ENTRY DATE: Entered STN: 19950307

Last Updated on STN: 19950307 Entered Medline: 19950222

AB OBJECTIVES. To study the cardiovascular effects of \*\*\*human\*\*\*

\*\*\*growth\*\*\* \*\*\*hormone\*\*\* (GH) \*\*\*replacement\*\*\*

in adults. INTERVENTION. Biosynthetic human GH given in a \*\*\*therapy\*\*\* \*\*\*daily\*\*\* \*\*\*dose\*\*\* of 0.04 +/- 0.01 IU kg-1 for 6-18 months in an open trial. PATIENTS. Thirty-four GH-deficient hypopituitary patients \*\*\*therapy\*\*\* , aged 19-67 years on conventional \*\*\*replacement\*\*\* and with a body mass index of 18.0-410.0 kg/m2. MEASUREMENTS. Resting blood pressure, exercise tolerance, renal function and routine blood counts were assessed every 6 months. Two-dimensional echocardiography and Doppler ultrasound scanning were performed at 0, 6 and 12 months of GH therapy. RESULTS. Exercise time increased significantly on GH from 9.37 +/- 2.64 min at the start to 10.39 +/- 2.86 min (P < 0.001), 10.90 +/-2.48 min (P < 0.001) and 11.11 + /- 0.70 min (P < 0.001) at 6, 12 and 18 months respectively. There was no change in the heart rate or in the blood pressure at rest nor at the peak of exercise. No significant changes were observed in measures of cardiac structure (left ventricular mass index, left ventricular posterior wall thickness and interventricular septal thickness), ejection fraction nor in cardiac output. Isovolumic relaxation time, a marker of diastolic function, decreased in 24 patients after 6

months on GH (from 98.6 +/- 15.9 to 89.6 +/- 15.2 ms; P < 0.031 but it was not different from baseline the 18 patients who were restured at 12 months. There was no significant change in the left ventricular filling neither at 6 nor at 12 months. No significant, changes were observed in plasma electrolytes, creatinine nor in blood count on GH treatment. CONCLUSIONS. Growth hormone \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* hypopituitary adults for 6-18 months produced sustained increase in exercise tolerance but was not associated with changes in cardiac structure or systolic function.

DUPLICATE 6 ANSWER 7 OF 12 MEDLINE

94236209 MEDLINE ACCESSION NUMBER:

DOCUMENT NUMBER: 94236209 PubMed ID: 8180671

Cardiovascular effects of growth hormone replacement TITLE:

therapy in hypopituitary adults.

**AUTHOR:** Beshyah S A; Shahi M; Skinner E; Sharp P; Foale R; Johnston

Unit of Metabolic Medicine, St Mary's Hospital and Medical CORPORATE SOURCE:

School, London, UK.

SOURCE: EUROPEAN JOURNAL OF ENDOCRINOLOGY, (1994 May) 130 (5)

451-8.

Journal code: 9423848. ISSN: 0804-4643.

PUB. COUNTRY: Norway

(CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199406

Entered STN: 19940621 ENTRY DATE:

> Last Updated on STN: 19940621 Entered Medline: 19940616

AR In the present study the effects of replacement with biosynthetic \*\*\*human\*\*\* hypopituitary adults on cardiac structure and function were investigated. Thirty-six GH-deficient, hypopituitary patients (17 males and 19 females; aged 19-67 years) on conventional \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* without GH were studied. Twenty-nine of the patients had acquired hypopituitarism in adult life, mainly due to pituitary tumours. The design of the study was a prospective, randomized, double-blind placebo-controlled trial for 6 months. Growth hormone (17 patients) was given in a \*\*\*daily\*\*\* \*\*\*dose\*\*\* of 0.02-0.05 IU/kg body wt sc (or a placebo, 19 patients) according to the patients' tolerance. Other pituitary replacement treatment was unchanged. Resting and exercise electrocardiography using the Bruce protocol, two-dimensional echocardiography, Doppler ultrasound scanning and serum insulin-like growth factor I (IGF-I) were assessed at 0 and 6 months. Resting blood pressure was measured at 0, 1, 3 and 6 months. Serum IGF-I increased significantly on GH treatment (mean +/- SD) GH: 293 +/- 197 vs placebo: 82 +/- 40 micrograms/l; p < 0.0001 at 6 months). Exercise time increased significantly on GH but not on placebo (GH: 8.45 +/- 3.16 to 9.38 +/- 2.42min.sec, p < 0.01; placebo 9.08 +/- 4.35 to 9.50 +/- 4.14 min.sec, NS), although the change was not significantly different between the two. There was no change in the heart rate or the blood pressure either at rest or at the peak of exercise. (ABSTRACT TRUNCATED AT 250 WORDS)

ANSWER 8 OF 12 MEDLINE DUPLICATE 7

ACCESSION NUMBER: 91193645 MEDLINE

PubMed ID: 2013748 DOCUMENT NUMBER: 91193645

TITLE: Evidence for the role of the secretory pattern of growth

hormone in the regulation of serum concentrations of

cholesterol and apolipoprotein E in rats.

AUTHOR: Oscarsson J; Carlsson L M; Bick T; Lidell A; Olofsson S O;

Eden S

CORPORATE SOURCE: Department of Physiology, University of Goteborg, Sweden. SOURCE:

JOURNAL OF ENDOCRINOLOGY, (1991 Mar) 128 (3) 433-8.

Journal code: 0375363. ISSN: 0022-0795.

PUB. COUNTRY: ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199105 ENTRY DATE:

Entered STN: 19910602 Last Updated STN: 3 STN: 19910602 Entered Medline: 19910515

Adult male Sprague-Dawley rats were hypophysectomized and connected to an AB automatic i.v. infusion system. The same \*\*\*daily\*\*\* \*\*\*dose\*\*\* human GH ( \*\*\*hGH\*\*\* ) was given either as eight \*\*\*daily\*\*\* (3-h intervals) to mimic the male specific secretory pattern of GH or as a continuous infusion of GH, to mimic the female secretory pattern. Hypophysectomized rats received i.v. \*\*\*replacement\*\*\* \*\*\*therapy\*\*\* with L-thyroxine and cortisol. The rats were treated for 5 days. The serum cholesterol concentration was higher when \*\*\*hGH\*\*\* was given continuously than when \*\*\*hGH\*\*\* was given as eight \*\*\*daily\*\*\* pulses. The concentration of high-density lipoprotein (HDL)-cholesterol was not influenced by intermittent GH treatment, but increased when was given as a continuous infusion. The serum concentration of apolipoprotein (Apo) E increased following treatment with a continuous infusion of \*\*\*hGH\*\*\* , whereas eight \*\*\*daily\*\*\* pulses of \*\*\*hGH\*\*\* had no effect. The serum concentration of ApoA-I was \*\*\*hGH\*\*\* treatment. The serum concentration of ApoB unaffected by decreased to the same degree whether \*\*\*hGH\*\*\* was given as a \*\*\*daily\*\*\* pulses. The serum continuous infusion or as eight concentration of triglycerides was not affected by \*\*\*hGH\*\*\* treatment. These results indicate that the higher serum HDL-cholesterol and serum ApoE concentrations of female rats may be due to their more continuous secretion of GH. In contrast, the effects of GH on the serum concentration of ApoB, which is not sexually differentiated, may be independent of the mode of GH secretion.

**DUPLICATE 8** ANSWER 9 OF 12 MEDLINE

ACCESSION NUMBER: 89383346 MEDLINE

DOCUMENT NUMBER: 89383346 PubMed ID: 2779233

Secretory pattern of growth hormone regulates steroid TITLE:

sulfatase activity in rat liver.

Eriksson L; Nilsson B; Carlstrom K; Oscarsson J; Eden S; AUTHOR:

von Schoultz B

Department of Obstetrics and Gynecology, University CORPORATE SOURCE:

Hospital, Umea, Sweden.

SOURCE: JOURNAL OF STEROID BIOCHEMISTRY, (1989 Sep) 33 (3) 413-6.

Journal code: 0260125. ISSN: 0022-4731.

ENGLAND: United Kingdom PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198910

ENTRY DATE: Entered STN: 19900309

Last Updated on STN: 19960129 Entered Medline: 19891019

AΒ Steroid sulfatase activity was quantified in liver microsomes from hypophysectomized adult female rats treated with estradiol and continuous \*\*\*hormone\*\*\* or intermittent \*\*\*human\*\*\* \*\*\*growth\*\*\* (

\*\*\*hGH\*\*\* ). Hypophysectomy clearly enhanced sulfatase activity as compared to intact female rats. Normal female values were completely restored by continuous infusion of \*\*\*hGH\*\*\* (1.4 i.u./kg/day). Neither the same \*\*\*dose\*\*\* of \*\*\*hGH\*\*\* given as two

\*\*\*daily\*\*\* injections nor estrogen \*\*\*replacement\*\*\*

had any effect. It is concluded that liver microsome \*\*\*therapy\*\*\* sulfatase activity in the non-pregnant rat is regulated by the sexually dimorphic secretory pattern of GH.

ANSWER 10 OF 12 MEDLINE

ACCESSION NUMBER: 87028782 MEDLINE

DOCUMENT NUMBER: 87028782 PubMed ID: 2429792

TITLE: Treatment of growth hormone deficiency.

AUTHOR: Ranke M B; Bierich J R

CLINICS IN ENDOCRINOLOGY AND METABOLISM, (1986 Aug) 15 (3) SOURCE:

495-510. Ref: 131

Journal code: 0357424. ISSN: 0300-595X.

PUB. COUNTRY: ENGLAND: United Kingdom

Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

LANGUAGE: English

FILE SEGMENT: Priority Journals ENTRY MONTH: 198612

ENTRY DATE: Entered STN: 00302

Last Updated on STN: 19900302

Entered Medline: 19861209

AB According to the results reported in the literature and from our own experience, the following recommendations for the treatment of children with GHD can be given: In order to start GH \*\*\*replacement\*\*\*

with GHD can be given: In order to start GH \*\*\*replacement\*\*\*

\*\*\*therapy\*\*\* in early childhood the diagnosis of GHD should be made as early as possible. The growth hormone \*\*\*dose\*\*\* during prepubertal age should not fall short of 12 IU/m2 per week. During spontaneous or

induced puberty, the \*\*\*dose\*\*\* needs to be increased, possibly by a factor of two. \*\*\*Daily\*\*\* subcutaneous injections appear most suitable. Treatment with growth hormone releasing factors in cases with hypothalamic GHD, although a promising alternative to the treatment with

\*\*\*hGH\*\*\* (Thorner et al, 1985), must be considered experimental at this point. Thyroxine replacement at a \*\*\*daily\*\*\* \*\*\*dose\*\*\* of 75-100 micrograms/m2 should be given in cases of secondary hypothyroidism. Glucocorticoid replacement, if required, should be given at low

\*\*\*doses\*\*\* (e.g. hydrocortisone 10 (to 15) mg/m2 per day in divided 
\*\*\*doses\*\*\* ). In cases with additional gonadotropin deficiency, sex 
steroids (or anabolic steroids) should be given with frequent monitoring 
of bone maturity not before the age of 13 in girls or 15 years in boys. In 
boys depot testosterone starting at low \*\*\*doses\*\*\* (e.g. 50-100 
mg/month i.m.) will induce a puberty-like increment in height velocity. 
Since the effect of oestrogens--even in low \*\*\*doses\*\*\* --on growth is 
uncertain, their administration before achievement of near-normal adult 
height should be avoided. With the advancement of diagnostic techniques 
and with the experience in treatment accumulated over the past 25 years, 
patients with GHD need no longer become dwarfs.

L6 ANSWER 11 OF 12 MEDLINE DUPLICATE 9

ACCESSION NUMBER: 83044754 MEDLINE

DOCUMENT NUMBER: 83044754 PubMed ID: 7136757

TITLE: Effect of frequency of growth hormone administration on

longitudinal bone growth and body weight in

hypophysectomized rats.

AUTHOR: Jansson J O; Albertsson-Wikland K; Eden S; Thorngren K G;

Isaksson O

SOURCE: ACTA PHYSIOLOGICA SCANDINAVICA, (1982 Feb) 114 (2) 261-5.

Journal code: 0370362. ISSN: 0001-6772.

PUB. COUNTRY: Sweden

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198212

ENTRY DATE: Entered STN: 19900317

Last Updated on STN: 19900317 Entered Medline: 19821216

AB The effect of frequency of growth hormone (GH) administration on longitudinal bone growth and body weight was studied in hypophysectomized rats. \*\*\*Replacement\*\*\* \*\*\*therapy\*\*\* with 3 different

\*\*\*doses\*\*\* of human GH [( \*\*\*hGH\*\*\* ) Crescormone] was started 10-14 days after hypophysectomy and was continued for 5 days. Longitudinal bone growth, as measured by the tetracycline method, and body weight were determined during the injection period. With a \*\*\*daily\*\*\* replacement \*\*\*dose\*\*\* of 128 micrograms of \*\*\*hGH\*\*\* body weight gain and

\*\*\*dose\*\*\* of 128 micrograms of \*\*\*nGH\*\*\* body Weight gain and longitudinal bone growth were significantly higher when the hormone was injected 4 and 8 times per day compared with animals receiving the hormone in one \*\*\*daily\*\*\* injection. When the \*\*\*dose\*\*\* of \*\*\*hGH\*\*\* was 32 or 8 micrograms per day, longitudinal bone growth and body weight gain were more pronounced in animals receiving the hormone 2 and 4 times per day compared with animals receiving the hormone one or 8 times per day. The results of the present study demonstrate that the frequency of GH administration influence body growth. The findings suggest that the secretory pattern of GH influence the growth rate under in vivo condition.

L6 ANSWER 12 OF 12 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 1982:158945 BIOSIS

DOCUMENT NUMBER: BA73:18929

TITLE: INTRA CAPSULAR IRRADIATION THERAPY OF CRANIO PHARYNGIOMAS

WITH RADIOACTIVE GOLD INDICATION AND FOLLOW-UP RESULTS.

AUTHOR(S): KODAMA T; MATSUKADO Y; UEMURA S

DEPARTMENT OF NEUROSURGERY, KUMAMOTO UNIVERSITY MEDICAL CORPORATE SOURCE: SCHOOL, KUMAM 860. NEUROL MED-CHIR, (1981) 21 (1), 49-58. SOURCE: CODEN: NMCHBN. ISSN: 0470-8105. BA; OLD FILE SEGMENT: English LANGUAGE: Of 27 patients with craniopharyngiomas, 16 were arbitrarily subjected to combined treatment of simple surgical evacuation and intracapsular irradiation with 198-Au. Follow-up studies were performed on 15 cases and they ranged from 6 mo. to 11 yr. One patient was omitted from the study because of a short postoperative period. Immediate postoperative morbidity and the endocrine functions at the end of the follow-up study were compared with those of the patient who underwent extensive surgical resection of the tumors. Intracapsular irradiation with 198-Au had satisfactory effects in the treatment of cystic craniopharyngioma, especially in recurrent cases of initially solid tumors, with respect to \*\*\*daily\*\*\* the preservation of the endocrine functions and the activity of the patients. The patients, who were followed up for over 5 yr, maintained an occupational IQ score in the normal range and the patients under school age were all able to continue their school lives. One of the female patients, who married after the treatment, was able to \*\*\*therapy\*\*\* have 2 children without any specific \*\*\*replacement\*\*\* . Another patient who had shown physical retardation in childhood due to \*\*\*growth\*\*\* \*\*\*hormone\*\*\* ] \*\*\*HGH\*\*\* [ \*\*\*human\*\*\* deficiency, showed favorable results with \*\*\*HGH\*\*\* administration in comparison with cases of extensive resection. Although the dosimetric value of 198-Au should be varied according to the size and thickness of the capsule, 15 to 30 mCi of 198-Au was the appropriate \*\*\*dose\*\*\* for treatment. => d his (FILE 'HOME' ENTERED AT 17:17:25 ON 02 JUL 2002) FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT 17:18:01 ON 02 JUL 2002 36627 S (HUMAN GROWTH HORMONE) OR HGH L1 92997 S (REPLACEMENT THERAPY) OR REPLENISH? L2456 S L1 (P) L2 L3122 S L3 (P) DOSE L437 S L4 (P) DAILY L5 12 DUPLICATE REMOVE L5 (25 DUPLICATES REMOVED) L6 => s microsphere 75372 MICROSPHERE L7 => s 11 (p) 17144 L1 (P) L7 L8 => s 18 (p) 12 L9 0 L8 (P) L2 => d his (FILE 'HOME' ENTERED AT 17:17:25 ON 02 JUL 2002) FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT 17:18:01 ON 02 JUL 2002 36627 S (HUMAN GROWTH HORMONE) OR HGH L192997 S (REPLACEMENT THERAPY) OR REPLENISH? L2456 S L1 (P) L2 L3L4122 S L3 (P) DOSE L5 37 S L4 (P) DAILY 12 DUPLICATE REMOVE L5 (25 DUPLICATES REMOVED) L6 L7 75372 S MICROSPHERE 144 S L1 (P) L7 L80 S L8 (P) L2 L9 => log y

=> log y COST IN U.S. DOLLARS STN INTERNATIONAL LOGOFF AT 17:22:28 ON 02 JUL 2002